Impact of social interactions in the community on the transmission of tuberculosis in a high incidence area

Collette N Classen, Robin Warren, Madeleine Richardson, John H Hauman, Robert P Gie, James H P Ellis, Paul D van Helden, Nulda Beyers

Abstract

Background—Tuberculosis (TB) is transmitted by close contact with an infectious person. It is assumed that close contact occurs amongst household members and that contact outside the house is “casual” and does not play a major role in the transmission of TB.

Methods—This study was conducted in an impoverished area with a high incidence of TB and a low HIV seropositive prevalence. Thirty three households with 84 TB patients were identified between February 1993 and April 1996 and the transmission of TB was studied by combining *Mycobacterium tuberculosis* fingerprinting with in depth sociological interviews.

Results—Forty two strain genotypes were identified in the 84 patients. In 15 households all the patients had identical strains, in nine households all the patients had different strains, and in nine households some patients had identical strains and one had a different strain. In 26 houses at least one patient had a strain which formed part of a larger community cluster and in 12 of these households the patient(s) had contact with a community member who had the identical strain. In 58% of the cases the contact took place while drinking in social groups.

Conclusion—In high incidence areas contact outside the household may be important for the transmission of TB. This contact often takes place during recreation which, in the case of this study of impoverished people, consisted of drinking in social groups. Social interaction patterns should be studied and understood for effective implementation of control strategies.

(Thorax 1999;54:136-140)

Keywords: tuberculosis; transmission; social interaction

drugs are regarded as risk factors for developing TB,⁶⁻⁸ but it is uncertain whether susceptibility to TB is increased by the alcohol and drugs per se, or by the fact that alcohol and drug users may be in close contact with untreated TB patients.

DNA fingerprinting of *M tuberculosis* strains by the insertion element IS6110 has recently indicated that, in patients with identical strains who are considered to reflect a chain of recent transmission, social contact could be proved only in a minority of cases.^{9,10} Additional probes^{11,12} classify *M tuberculosis* more accurately into clusters, thereby increasing the likelihood of finding contact between patients.¹¹ Limited variation has been recorded in *M tuberculosis* strains from households¹³ and close contacts,¹⁴ suggesting transmission of *M tuberculosis*. However, in these studies the possibility that the transmission might have taken place in the community was not addressed.

The aim of this study was to investigate the transmission of TB in households by using IS6110 and MTB484(1) DNA fingerprinting of *M tuberculosis* organisms from patients in the same household, combined with an in depth anthropological assessment of social interaction patterns in a high incidence area.

Methods

The study was performed between February 1993 and April 1996 in the Western Cape Province of South Africa where the tuberculosis notification rate for 1995 was 682/100 000¹⁵ and the prevalence of HIV seropositivity is just more than 1%.¹⁶ The tuberculosis notification rate in the suburbs studied (population 34 294, surface area 2.42 km²) was >1000/100 000.¹⁷ Treatment for TB is provided by two local authority health clinics and patients receive a six month antituberculosis treatment regimen daily under direct supervision (DOTs).

During the study period all sputum samples of TB patients were cultured by radiometric assay (BACTEC[®]) and subcultured onto Löwenstein-Jensen slants. DNA was isolated and genotyped by IS6110 and MTB484(1).^{11,18} The IS6110 DNA fingerprints were analysed using GelCompar software and compared using the UPGMA and Dice coefficients.¹⁹ The MTB484(1) DNA fingerprints were analysed visually.

M tuberculosis strains were classified into clusters if two or more people had an identical strain with both probes. If a number of people in different households had the identical strain

Department of
Anthropology/Sociology,
University of the
Western Cape, South
Africa
C N Classen
J H P Ellis

Department of
Paediatrics & Child
Health, University of
Stellenbosch, South
Africa
R P Gie
N Beyers

MRC Centre for
Molecular and Cellular
Biology, Department
of Medical
Biochemistry,
University of
Stellenbosch, South
Africa
R Warren
M Richardson
J H Hauman
P D van Helden

Correspondence to:
Dr C Classen, Department
of Paediatrics and Child
Health, University of
Stellenbosch, PO Box 19063,
Tygerberg 7505, South
Africa.

Received 29 July 1998
Returned to author
23 September 1998
Revised manuscript received
10 November 1998
Accepted for publication
11 November 1998

Transmission of *Mycobacterium tuberculosis* occurs due to contact with an infectious individual¹ and several clinical and epidemiological studies of infection or disease after contact with tuberculosis (TB) have been published.^{2,3} Although households are regarded as places of close contact and high risk for transmission of TB, the exact definition of a “household” or “close contact” is not clear^{4,5} and the relative importance of crowding or contact with a TB patient outside the household has not been established.^{2,5,6} Alcohol and

it was called a “community cluster”. If all the people with an identical strain lived in the same household the strain was called a “unique household cluster”. If a strain did not match any other strain from the community it was called a “unique strain”.

Information regarding contact between patients was obtained by a medical anthropologist through informal interviews conducted in patients' homes or in a secluded part of the clinic. Occasionally patients were met in neighbourhood bars in order to observe the alcohol consumption and the social contact in these bars. Confidentiality was maintained at all times.

We defined a household as a house and associated informal dwelling(s) at the same address on the same plot of land. Plot sizes ranged in size from 250 m² to 800 m², there was always a single entrance to the plot and access to the associated informal dwellings was obtained by walking through the main house.

Results

IS6110 AND MTB484(1) FINGERPRINTING AND SOCIAL INTERVIEWS

During the study period (February 1993 to April 1996) 622 patients residing in the area were shown to be infected with *M. tuberculosis* organisms, having 283 different banding patterns on IS6110 fingerprinting, and the degree of clustering was calculated at 40–50%. During

this period 44 of the 5345 households in the community were identified in which two or more household members had cultures positive for *M. tuberculosis*. Eleven households were excluded from the study as all the household members could not be interviewed (in one household both patients had died while at least one patient of each of the remaining 10 households had moved). The final sample consisted of 33 households with 84 patients who were all interviewed and whose cultures revealed 42 *M. tuberculosis* strains. There were 26 households (79%) in which at least one patient had a strain which formed part of a community cluster. In five of the 33 households there was a strain of *M. tuberculosis* shared by the patients in that specific household but which did not occur in any other member of the community (unique household clusters). Of the 84 patients 14 (17%) had a unique *M. tuberculosis* strain not shared by any other patient in the community.

The households were grouped into three different categories.

(1) Households where patients in each house had identical strains

In all 15 households where the patients within the same household had identical *M. tuberculosis* strains the patients in the same household shared the same dwelling (table 1) and in 14 households the patients were close relatives. In

Table 1 Data of patients and *Mycobacterium tuberculosis* strains in households where all patients per household had the identical strain of *M. tuberculosis*

Household	No. of bands with IS6110	No. of patients with identical strains	Relationship	UC/CC	No. of other people in community cluster	No. of household patients drinking in social group	No. of other people in community cluster in contact with patient(s)
1	11	2*	Uncle and niece	CC	2	1	0
2	19	4*	Brothers and sister, daughter	CC	7	3	1
3	4	2*	Common-law spouses	CC	15	2	3(s)
4	3	2*	Ex-spouses	UC	-	1	0
5	3	2*	Brothers	CC	3	2	0
6	1	2*	Friends	CC	6	2	1(s)
7	4	2*	Mother and son	CC	4	1	0
8	9	2*	Father and son	CC	2	2	0
9	14	2*	Father and son	UC	-	1	0
10	9	3*	Mother, daughter and cousin	UC	-	1	0
11	14	2*	Spouses	CC	3	1	0
12	17	2*	Mother and son	CC	1	2	0
13	5	2*	Common-law spouses	CC	8	2	1
14	14	2*	Brother and cousin	CC	2	2	1(s)
15	14	6*	Brothers, sister and sister's boyfriend	CC	10	6	0

UC = unique household cluster; CC = community cluster; *same dwelling; (s) = social contact took place while drinking.

Table 2 Data of patients and *Mycobacterium tuberculosis* strains in households where each patient in a household had a different strain of *M. tuberculosis*

Household	No. of bands with IS6110	No. of patients with identical strains	Relationship	U/UC/CC	No. of other people in cluster	No. of household patients drinking in social group	No. of other people in community cluster in contact with patient(s)
1	12	2*	Uncle and niece	U	-	1	-
	4			CC	6		1
2	10	2*	Tenant	CC	1	2	0
15	15			U	-		-
3	12	2†	Tenant	U	-	1	-
14	14			U	-		-
4	11	2†	Tenant	CC	4	2	3(s)
21	21			CC	1		0
5	10	2*	Tenant	U	-	2	-
12	12			CC	1		1
6	9	2†	Cousins	CC	10	2	0
12	12			CC	2		0
7	9	2*	Spouses	U	-	2	-
14	14			U	-		-
8	14	2*	Tenant	CC	3	2	0
15	15			CC	5		2(s)
9	15	2†	Tenant	U	-	2	-
4	4			U	-		-

U = unique strain; UC = unique household cluster; CC = community cluster; *same dwelling; †different dwelling; (s) = social contact took place while drinking.

Table 3 Data of patients and *Mycobacterium tuberculosis* strains in households where two or more patients had the identical *M tuberculosis* strain and one patient had a different strain

Household	No. of bands with IS6110	No. of patients with identical strains	Relationship	UC/CC	No. of other people in community cluster	No. of household patients drinking in social group	No. of other people in community cluster with patient(s)	No. of patients with different strains	No. of bands with IS6110 strain typing	Relationship to patients with same strain	UC/CC**	No. of patients in cluster with same strain	No. of household patients drinking in social groups	No. of other people in community cluster in contact with patient(s)
1	13	2	Family*	CC	1	2	0	1	12	Acquaintances*	U	-	1	-
2	15	2	Common-law spouses*	CC	4	2	1(s)	1	11	Tenant†	CC	5	1	No
3	5	2	Brother and sister*	CC	1	1	0	1	16	Sister*	U	-	1	-
4	11	2	Grandfather and granddaughter*	CC	6	1	1	1	11	Tenant*	U	-	1	-
5	11	2	Mother and son*	UC	-	1	-	1	15	Husband and father*	U	-	1	-
6	12	2	Acquaintances*	CC	-	2	0	1	13	Acquaintances*	CC	1	1	No
7	13	2	Common-law spouses*	UC	-	2	-	1	5	Tenant*	CC	1	1	No
8	4	3	Acquaintances*	CC	14	3	2(s)	1	4	Acquaintances*	U	-	1	-
9	11	2	In-laws†	CC	4	2	0	1	19	Tenant†	CC	10	1	No

UC = unique household cluster; CC = community cluster; U = unique strain; *same dwelling; †different dwelling; (s) = social contact took place while drinking.

three of the households the patients in the household had identical *M tuberculosis* organisms which formed a unique household cluster and the organism did not form part of a bigger cluster shared by other patients in the community. In 12 households the *M tuberculosis* strain formed part of a bigger community cluster (on average six other people in the cluster). At least one patient per household regularly drank in social groups with community members who had tuberculosis. In patients from five households, close contact with a community member with an identical *M tuberculosis* strain could be confirmed and in patients from three of the five households (60%) this contact occurred while drinking in social groups.

(2) Households where each patient in the household had a different strain

Of the 18 patients living in the nine households where patients in each household had different *M tuberculosis* strains, nine had unique strains and nine had strains which formed part of a community cluster (on average four other patients in the cluster; table 2). Regular drinking in social groups occurred in 16 of the 18 patients. In four patients whose strain formed part of a community cluster, contact with other community members with the identical *M tuberculosis* strain could be proved, and in two of the four cases this contact took place while drinking in social groups. In only two of the nine households were the patients close relatives or spouses, and in six of the nine households the patients lived in the same dwelling.

(3) Households where at least two patients had identical strains and at least one patient had a different strain

There were nine households with a total of 28 patients where two or more patients with TB in the same household had identical *M tuberculosis* strains and an additional patient had a strain different from the others (table 3). There were 19 patients in eight of the households who had an *M tuberculosis* strain which formed part of a community cluster and 17 of these regularly drank in social groups. In three cases close contact with other patients in the community who had the identical *M tuberculosis* strain was identified and in two of the three cases the contact occurred while drinking in social groups. In all the households the patients who had identical *M tuberculosis* strains lived in the same dwelling and in six of the nine dwellings these patients were relatives. In three of the households the patient with the different *M tuberculosis* strain lived in a different dwelling on the same plot of land.

DRINKING IN SOCIAL GROUPS

Social contact outside the household was established in members of 12 of the 26 households where the *M tuberculosis* strains in the household formed part of a community cluster. In seven of these 12 households (58%) this contact took place during drinking in social groups. Of the 84 patients in this household study, 62 (74%) regularly drank in social

groups. This type of drinking took place either in a local neighbourhood informal bar (shebeen) or in small gatherings at peoples' homes. In the shebeens 30–40 people drank together in a space of about 35 m², while in the homes usually 3–6 people gathered in a 9–10 m² area while drinking. Each patient drank 4–5 times per week and would often spend up to 30 hours a week drinking in social groups.

Discussion

There have been numerous reports of TB and of skin test conversion amongst family members,^{2,5} suggesting transmission in the house. Recently two reports of family members or closely associated individuals living in the same house^{13,14} indicated limited variation in the RFLP patterns, suggesting transmission in the house. No indication is given on their possible contact with other TB patients outside the house.

In communities where people live in overcrowded conditions one would expect that, if two or more people in one household have TB, their *M. tuberculosis* strains would be identical. Our findings from an area with a high incidence of TB demonstrate that, when two or more people in the same household have TB, they do not always have the same strain of *M. tuberculosis* and therefore household transmission cannot always be implied. Transmission as defined by identical *M. tuberculosis* strains occurred more commonly in patients who were relatives living in the same dwelling than in non-related individuals or if relatives lived in different dwellings. We have found that, in 18 of the 33 (55%) households, at least one patient had a strain which was not shared by another household member, suggesting that tuberculosis is being transmitted either outside the households or occurs due to reactivation of latent infection. In seven of these households the different strain could be explained by the fact that one patient of each household lived in a separate dwelling (backyard shack). In the community studied, where there is a high incidence of TB (>1000/100 000/year) and where people live in conditions of poverty, the difficulty of tracing the transmission of TB is illustrated by the complexity of the household structures where living at the same address does not always imply living in the same dwelling. Even when people live in the same dwelling they often do not even know each other as their living arrangements are merely business transactions and do not necessarily imply close contact with each other.

Our study suggests that, even when household members have identical *M. tuberculosis* strains, transmission within the household cannot be assumed since the *M. tuberculosis* strain in 59 of the 84 patients (70%) in these households formed part of a community cluster. In 12 of the households contact with non-household members with the identical *M. tuberculosis* strain had taken place and in seven of these 12 (58%) households contact took place while drinking in social groups. During these regular drinking episodes very close con-

tact (kissing, hugging, dancing) occurred in confined spaces (1–2 m² per person).

Contact outside the house or the family including casual, short periods of contact amongst family members and co-workers or casual contact in a physician's office,²¹ in shelters for the urban homeless,^{22–24} housing facilities for HIV positive patients,²⁵ a church choir,²⁶ a church,²⁷ and in a crack house²⁸ has been reported as important in the transmission of TB. A neighbourhood bar has been implicated in the transmission of TB amongst its regular patrons, but without RFLP evidence.²⁹ Transmission of TB in bars has been proved by RFLP data, but only in HIV infected individuals.³⁰ These studies all looked at the transmission of TB within relatively confined areas and did not report the effect of transmission of TB in the wider community outside these confined areas.

We therefore suggest that, while in low incidence areas transmission of TB often takes place in confined areas like the household, in high incidence areas contact outside the house may play a major role in the transmission of TB. In our study, in an economically depressed area with no recreation except for social drinking, we found that 74% of all the patients drank regularly in social groups. Our findings further indicate that contact outside the house is not necessarily "casual" and that, while drinking in social groups, very close and intimate contact occurred which could easily have contributed to the transmission of TB. Similar conditions are probably prevailing in many other developing countries with high incidences of TB, thus making extrapolation from studies on the transmission of TB in low incidence developed countries impossible and dangerous. We suggest that the social interaction patterns in a community should be studied before the transmission of TB in the area can be properly understood and effective TB control strategies can be put into place.

We are indebted to the people of Uitsig and Ravensmead who effortlessly allowed us a glimpse into very personal spheres of their lives, Mrs S M Carlini and Mrs M Bosman for help with the *M. tuberculosis* cultures, Dr I Toms and Dr T Fish for permission to conduct the study in the local authority health clinics, Glaxo Wellcome Action TB Research Program for financial support, and the Commission of the European Communities, Directorate General XII, Biomedical and Health Research, Biomed 1, contract BMH1-CT93-1614 for assistance with the Gelcomp software.

- Coovadia HM, Benatar SR. *A century of tuberculosis*. Cape Town: Oxford University Press, 1991.
- Chapman JS, Dyerly MD. Social and other factors in intra-familial transmission of tuberculosis. *Am Rev Respir Dis* 1964;90:48–60.
- Narain R, Nair SS, Rao GR, et al. Distribution of tuberculous infection and disease among households in a rural community. *Bull World Health Org* 1966;34:639–54.
- Lomnitz L. *Networks and marginality. Life in a Mexican shanty town*. New York: Academic Press, 1977.
- Hussain SF, Watura R, Cashman B, et al. Tuberculosis contact tracing: are the British Thoracic Society guidelines still appropriate? *Thorax* 1992;47:984–5.
- Coetzee N, Yach D, Joubert G. Crowding and alcohol abuse as risk factors for tuberculosis in the Mamre population. *S Afr Med J* 1988;74:352–4.
- Enarson DA, Wang J, Dirks JM. The incidence of active tuberculosis in a large urban area. *Am J Epidemiol* 1989;129:1268–76.
- Bloch AB, Onorato IM, Ihle WW, et al. The need for epidemic intelligence. *Public Health Rep* 1996;111:26–31.
- Alland D, Kalkut G E, Moss AR, et al. Transmission of tuberculosis in New York City: an analysis by DNA fingerprinting and conventional epidemiologic methods. *N Engl J Med* 1994;330:1710–16.

- 10 Small PM, Hopewell PC, Sing SP, et al. The epidemiology of tuberculosis in San Francisco: population-based study using conventional and molecular methods. *N Engl J Med* 1994;330:1703-9.
- 11 Warren RM, Richardson M, Sampson S, et al. Genotyping of *Mycobacterium tuberculosis* with additional markers enhances accuracy in epidemiological studies. *J Clin Microbiol* 1996;34:2219-24.
- 12 Burman WJ, Reeves RR, Hawkes AP, et al. DNA fingerprinting with two probes decreases clustering of *Mycobacterium tuberculosis*. *J Clin Microbiol* 1996;34:2219-24.
- 13 Mazurek GH, Cave MD, Eisenach KD, et al. Chromosomal DNA fingerprint patterns produced with IS6110 as strain-specific markers for epidemiologic study of tuberculosis. *J Clin Microbiol* 1991;29:2030-3.
- 14 Huh YJ, Ahn DI, Kim SJ. Limited variation of DNA fingerprints (IS6110 and IS1081) in Korean strains of *Mycobacterium tuberculosis*. *Tubercle Lung Dis* 1995;76:324-9.
- 15 Department of Health, Republic of South Africa. Notifiable medical conditions. *Epidemiological Comments* 1996;23:22.
- 16 Department of Health, Republic of South Africa. Sixth national HIV survey of women attending antenatal clinics of the public health services in the Republic of South Africa, October/November 1995. *Epidemiological Comments* 1996;33(1):14.
- 17 Beyers N, Gie RP, Zietsman HL, et al. The use of geographical information system (GIS) to evaluate the distribution of tuberculosis in a high incidence community. *S Afr Med J* 1996;86:40-4.
- 18 Warren R, Hauman J, Beyers N, et al. Unexpectedly high strain diversity of *Mycobacterium tuberculosis* in a high incidence community. *S Afr Med J* 1996;86:45-9.
- 19 Hermans PWM, Messadi F, Guebrevabher H, et al. Analysis of the population structure of *Mycobacterium tuberculosis* in Ethiopia, Tunisia and The Netherlands. Usefulness of DNA typing for global epidemiology. *J Infect Dis* 1995;171:1504-13.
- 20 Festenstein F. Spread of tuberculosis within a family. *Lancet* 1981;i:603-5.
- 21 Valway SE, Sanchez MPC, Shinnick TF, et al. An outbreak involving extensive transmission of a virulent strain of *Mycobacterium tuberculosis*. *N Engl J Med* 1998;338:633-9.
- 22 Barnes PF, El-Hajj H, Preston-Martin S, et al. Transmission of tuberculosis among the urban homeless. *JAMA* 1996;274:305-7.
- 23 Barnes PF, Yang Z, Preston-Martin S, et al. Patterns of tuberculosis transmission in central Los Angeles. *JAMA* 1997;278:1159-63.
- 24 Gutierrez MC, Vincent V, Aubert D, et al. Molecular fingerprinting of *Mycobacterium tuberculosis* and risk factors for tuberculosis transmission in Paris, France, and surrounding area. *J Clin Microbiol* 1998;36:486-92.
- 25 Daley CL, Small PM, Schecter GF, et al. An outbreak of tuberculosis with accelerated progression among persons infected with the Human Immunodeficiency Virus. *N Engl J Med* 1992;326:231-5.
- 26 Mangura BT, Napolitano EC, Passannante MR, et al. *Mycobacterium tuberculosis* miniepidemic in a church gospel choir. *Chest* 1998;113:234-7.
- 27 Dutt AK, Mehta JB, Whitaker BJ, et al. Outbreak of tuberculosis in a church. *Chest* 1995;107:447-52.
- 28 Leonhardt KK, Gentile F, Gilbert BP, et al. A cluster of tuberculosis among crack house contacts in San Mateo County, California. *Am J Public Health* 1994;84:1834-6.
- 29 Kline SE, Hedemark LL, Davies SF. Outbreak of tuberculosis among regular patrons of a neighborhood bar. *N Engl J Med* 1995;333:222-7.
- 30 Tabet SR, Goldbaum GM, Hooton TM, et al. Restriction fragment length polymorphism analysis detecting a community-based tuberculosis outbreak among persons infected with human immunodeficiency virus. *J Infect Dis* 1994;169:189-92.